

**Amendments to the claims:**

This listing of claims will replace all prior versions, and listings, of claims in the application:

**Listing of claims:**

Claims 1-56 (canceled)

57. (Previously presented) A method for characterizing samples having fluorescent particles, comprising the steps of:

- a) monitoring intensity fluctuations of fluorescence emitted by the particles in at least one measurement volume by detecting sequences of photon counts by at least one photon detector,
- b) determining from the sequences of photon counts intermediate statistical data comprising at least two probability functions,  $\hat{P}_1(\mathbf{n}_1), \hat{P}_2(\mathbf{n}_2), \dots$ , of the number of photon counts,  $\mathbf{n}_1, \mathbf{n}_2, \dots$ , detected in different sets of counting time intervals,
- c) determining from said intermediate statistical data a distribution of particles as a function of at least two arguments, wherein one argument is a specific brightness of the particles, or a measure thereof, and another argument is a diffusion coefficient of the particles, or a measure thereof,

wherein said distribution function of particles is determined by fitting the experimentally determined probability functions  $\hat{P}_1(\mathbf{n}_1), \hat{P}_2(\mathbf{n}_2), \dots$  by corresponding theoretical probability functions  $P_1(\mathbf{n}_1), P_2(\mathbf{n}_2), \dots$

and

wherein said theoretical probability distributions  $P_1(\mathbf{n}_1), P_2(\mathbf{n}_2), \dots$  are calculated as functions of apparent concentrations and apparent brightness which depend on the widths of the counting time intervals in the different sets.

58. (Previously presented) The method according to claim 57 wherein, in calculations of the theoretical distributions  $P_1(\mathbf{n}_1), P_2(\mathbf{n}_2), \dots$ , an optical spatial brightness function  $B(\mathbf{r})$  is accounted for by a separately determined relationship between brightness  $B$  and volume elements  $dV$ .
59. (Previously presented) The method according to claim 58, wherein the relationship between the spatial brightness  $B$  and volume elements  $dV$  is expressed through a variable  $x = \ln(B_0 / B)$  by a relationship  $\frac{dV}{dx} = A_0(1 + a_1x + a_2x^2)x^{a_3}$ , where  $B_0$  is maximum brightness and  $A_0, a_1, a_2$  and  $a_3$  are empirical parameters of the spatial brightness function.
60. (Previously presented) The method according to claim 57 wherein each set of counting time intervals consists of intervals of equal width while different probability functions  $\hat{P}_{T_1}(\mathbf{n}_1), \hat{P}_{T_2}(\mathbf{n}_2), \dots$  correspond to counting time intervals of different widths  $T_1, T_2, \dots$ .

61. (Previously presented) The method according to claim 60, wherein the apparent concentration is calculated as

$$c_{app}(T) = \frac{c_{app}(0)}{\Gamma(T)},$$

the apparent brightness is calculated as

$$q_{app}(T) = q_{app}(0)\Gamma(T),$$

and  $\Gamma(T)$  is calculated as

$$\Gamma(T) = \frac{1}{c_{app}(0)q_{app}^2(0)T^2} \int_0^T dt_1 \int_0^T dt_2 G(t_2 - t_1),$$

where  $G(t)$  denotes autocorrelation function of fluorescence intensity and  $T$  denotes the width of the counting time interval.

62. (Currently amended) The method according to claim 57 wherein the counting time intervals in each set are consecutive in time.

63. (Previously presented) The method according to claim 57 wherein, counting time intervals in each set overlap.

64. (Previously presented) The method according to claim 57 wherein said intermediate statistical data are processed applying inverse transformation with regularization and/or constraints.

65. (Previously presented) The method according to claim 57 wherein the theoretical distributions  $P_1(\mathbf{n}_1), P_2(\mathbf{n}_2), \dots$  are calculated through their generating functions

$$G_{P(\mathbf{n})}(\vec{\xi}) = \sum_n \vec{\xi}^n P(\mathbf{n}).$$

66. (Previously presented) The method according to claim 57 wherein said distribution function of particles is determined by fitting the experimentally determined probability functions  $\hat{P}_1(n_1), \hat{P}_2(n_2), \dots$  by corresponding theoretical probability functions  $P_1(n_1), P_2(n_2), \dots$ .

67. (Currently amended) The method according to claim 63 66 wherein the theoretical probability functions  $P_1(n_1), P_2(n_2), \dots$  are calculated through their generating functions

$$G_{P(n)}(\xi) = \sum_n \xi^n P(n).$$

68. (Currently amended) The method according to claim 62 65, wherein the generating function

is calculated using the expression  $G(\xi) = \exp[\int dq c(q) \int d^3 \mathbf{r} (e^{(\xi-1)qTB(\mathbf{r})} - 1)]$ ,

where  $c(q)$  is the apparent density of particles with specific brightness  $q$ ,  $T$  is the length of the counting time interval, and  $B(\mathbf{r})$  is the spatial brightness profile as a function of coordinates.

69. (Currently amended) The method according to claim 63 65, wherein said generating

functions are calculated using the formula  $G_{P(n)}(\xi) = \exp[\sum_i c_i \int (e^{(\xi-1)q_i B(\mathbf{r})T} - 1) dV]$  in

which  $c$  is an apparent concentration and  $q$  is an apparent brightness which both depend on the width of the counting time interval  $T$ .

70. (Previously presented) The method according to claim 57 wherein concentrations of particles are selected to be approximately one or less molecules per measurement volume.

71. (Currently amended) The method according to claim 57 wherein said at least one photon detector is either an avalanche photodiode or a photomultiplier.

72. (Currently amended) The method according to claim 57 wherein at least one photon

detectors are at least two photon detectors which are used for monitoring fluorescence of different wavelengths or polarization.

73. (Previously presented) The method according to claim 57 wherein said fluorescent particles are characterized by applying an homogeneous fluorescence assay.
74. (Currently amended) The method according to claim 57, wherein the intensity fluctuation of fluorescence emitted by the particles in said at least one measurement volume are monitored by detecting said sequences of photon counts using said at least one photon detector, said at least one which photon detector being is part of a confocal apparatus.
75. (Currently amended) The method according to claim 74, wherein said confocal apparatus comprises:
  - a) a radiation source (12) for providing excitation radiation (14),
  - b) an objective (22) for focusing the excitation radiation (14) into a measurement volume (26) of said at least one measurement volume,
  - c) a detector (42) of said at least one photon detector for detecting emission radiation (30) that stems from the measurement volume (26), and
  - d) an opaque means (44) positioned in the pathway (32) of the emission radiation (30) or excitation radiation (14) for erasing the central part of the emission radiation (30) or excitation radiation (14).

**Amendments to the Drawings:**

Six (6) replacement (corrected) sheets of drawings are attached hereto. These sheets contain Fig. 1 - Fig. 7, corrected pursuant to the PTO Notice (Form PTO-948) dated January 13, 2004 (copy attached).

Attachments: Replacement Sheets (6)  
Copy of PTO Notice